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RAW SEQUENCE LISTING DATE: 08/31/2004
PATENT APPLICATION: US/10/817,334 TIME: 15:03:51

Input Set : A:\-1310-1.app

Output Set: N:\CRF4\08312004\J817334.raw

3 <110 > APPLICANT: Hammock, Bruce D. 4 Kim, In-Hae 5 Morisseau, Christophe Watanabe, Takaho Newman, John W. The Regents of the University of California 10 <120> TITLE OF INVENTION: Improved Inhibitors for the Soluble Epoxide Hydrolase 12 <130> FILE REFERENCE: 02307W-131010US 14 <140> CURRENT APPLICATION NUMBER: US 10/817,334 15 <141> CURRENT FILING DATE: 2004-04-02 17 <150> PRIOR APPLICATION NUMBER: US 60/460,559 18 <151> PRIOR FILING DATE: 2003-04-03 20 <160> NUMBER OF SEQ ID NOS: 4 22 <170> SOFTWARE: PatentIn Ver. 2.1 24 <210> SEQ ID NO: 1 25 <211> LENGTH: 555 26 <212> TYPE: PRT 27 <213> ORGANISM: Homo sapiens 29 <220> FEATURE: 30 <223> OTHER INFORMATION: human soluble epoxide hydrolase (sEH) 32 <400> SEQUENCE: 1 33 Met Thr Leu Arg Ala Ala Val Phe Asp Leu Asp Gly Val Leu Ala Leu 34 1 36 Pro Ala Val Phe Gly Val Leu Gly Arg Thr Glu Glu Ala Leu Ala Leu 37 39 Pro Arg Gly Leu Leu Asn Asp Ala Phe Gln Lys Gly Gly Pro Glu Gly 35 40 42 Ala Thr Thr Arg Leu Met Lys Gly Glu Ile Thr Leu Ser Gln Trp Ile 55 45 Pro Leu Met Glu Glu Asn Cys Arg Lys Cys Ser Glu Thr Ala Lys Val 70 75 48 Cys Leu Pro Lys Asn Phe Ser Ile Lys Glu Ile Phe Asp Lys Ala Ile 51 Ser Ala Arg Lys Ile Asn Arg Pro Met Leu Gln Ala Ala Leu Met Leu 52 100 105 54 Arg Lys Lys Gly Phe Thr Thr Ala Ile Leu Thr Asn Thr Trp Leu Asp 55 115 120 57 Asp Arg Ala Glu Arg Asp Gly Leu Ala Gln Leu Met Cys Glu Leu Lys 135 60 Met His Phe Asp Phe Leu Ile Glu Ser Cys Gln Val Gly Met Val Lys 150 155 63 Pro Glu Pro Gln Ile Tyr Lys Phe Leu Leu Asp Thr Leu Lys Ala Ser 165 170

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	Ala	Arg	Asp		Gly	Met	Val	Thr 200		Leu	Val	Gln	Asp	190 Thr	Asp	Thr
	Ala	Leu 210	Lys	Glu	Leu	Glu	Lys 215		Thr	Gly	Ile	Gln 220		Leu	Asn	Thr
	Pro 225		Pro	Leu	Pro	Thr 230		Cys	Asn	Pro	Ser 235		Met	Ser	His	Gly 240
78 79	Tyr	Val	Thr	Val	Lys 245	Pro	Arg	Val	Arg	Leu 250		Phe	Val	Glu	Leu 255	
81 82	Ser	Gly	Pro	Ala 260	Val	Cys	Leu	Cys	His 265	Gly	Phe	Pro	Glu	Ser 270		\mathtt{Tyr}
84 85	Ser	Trp	Arg 275	Tyr	Gln	Ile	Pro	Ala 280	Leu	Ala	Gln	Ala	Gly 285	Tyr	Arg	Val
87 88	Leu	Ala 290	Met	Asp	Met		Gly 295	Tyr	Gly	Glu		Ser 300	Ala	Pro	Pro	Glu
91	305		Glu			310					315					320
94			Lys		325					330			_		335	_
97				340					345					350		
100			Val 355					360					365			
103		370			Glu	Ser	Ile 375		Ala	Asn	Pro	Val 380		Asp	Tyr	Gln
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106	385					390					395	Glu	Leu			Asn 400
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106 108 109 111 112 114 115	385 Leu Leu Pro	Ser Ser Glu	Arg Met Glu 435	Thr His 420 Pro	Phe 405 Lys Ser	390 Lys Val Leu	Ser Cys Ser	Leu Glu Arg 440	Phe Ala 425 Met	Arg 410 Gly Val	395 Ala Gly Thr	Glu Ser Leu Glu	Leu Asp Phe Glu 445	Glu Val 430 Glu	Ser 415 Asn	400 Val Ser Gln
106 108 109 111 112 114 115 117	Jeu Leu Pro	Ser Glu Tyr 450	Arg Met Glu 435 Val	Thr His 420 Pro	Phe 405 Lys Ser	390 Lys Val Leu Phe	Ser Cys Ser Lys 455	Leu Glu Arg 440 Lys	Phe Ala 425 Met Ser	Arg 410 Gly Val Gly	395 Ala Gly Thr	Ser Leu Glu Arg 460	Asp Phe Glu 445 Gly	Glu Val 430 Glu Pro	Ser 415 Asn Ile	400 Val Ser Gln Asn
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106 108 109 111 112 114 115 117 118 120 121 123 124	Leu Pro Phe Trp 465 Gly	Ser Ser Glu Tyr 450 Tyr	Arg Met Glu 435 Val Arg Lys	Thr His 420 Pro Gln Asn	Phe 405 Lys Ser Gln Met Leu 485	January Val Leu Phe Glu 470 Ile	Ser Cys Ser Lys 455 Arg	Leu Glu Arg 440 Lys Asn Ala	Phe Ala 425 Met Ser Trp	Arg 410 Gly Val Gly Lys Met 490	395 Ala Gly Thr Phe Trp 475 Val	Ser Leu Glu Arg 460 Ala Thr	Asp Phe Glu 445 Gly Cys Ala	Glu Val 430 Glu Pro Lys Glu	Ser 415 Asn Ile Leu Ser Lys 495	400 Val Ser Gln Asn Leu 480 Asp
106 108 109 111 112 114 115 117 118 120 121 123 124 126 127	Leu Pro Phe Trp 465 Gly Phe	Ser Ser Glu Tyr 450 Tyr Arg Val	Arg Met Glu 435 Val Arg Lys Leu	Thr His 420 Pro Gln Asn Ile Val 500	Phe 405 Lys Ser Gln Met Leu 485 Pro	January Val Leu Phe Glu 470 Ile Gln	Ser Cys Ser Lys 455 Arg Pro	Leu Glu Arg 440 Lys Asn Ala Ser	Phe Ala 425 Met Ser Trp Leu Gln 505	Arg 410 Gly Val Gly Lys Met 490 His	395 Ala Gly Thr Phe Trp 475 Val	Ser Leu Glu Arg 460 Ala Thr	Asp Phe Glu 445 Gly Cys Ala Asp	Glu Val 430 Glu Pro Lys Glu Trp 510	Ser 415 Asn Ile Leu Ser Lys 495 Ile	400 Val Ser Gln Asn Leu 480 Asp
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106 108 109 111 112 114 115 117 118 120 121 123 124 126 127 129 130 132 133 135 136	James Asp	Ser Ser Glu Tyr 450 Tyr Arg Val Leu Lys 530 Ala	Arg Met Glu 435 Val Arg Lys Leu Lys 515	Thr His 420 Pro Gln Asn Ile Val 500 Arg Thr	Phe 405 Lys Ser Gln Met Leu 485 Pro Gly Glu Pro	January Val Leu Phe Glu 470 Ile Gln His Val	Ser Cys Ser Lys 455 Arg Pro Met Ile Asn 535	Leu Glu Arg 440 Lys Asn Ala Ser Glu 520 Gln	Phe Ala 425 Met Ser Trp Leu Gln 505 Asp	Arg 410 Gly Val Gly Lys Met 490 His Cys	395 Ala Gly Thr Phe Trp 475 Val Met Gly Ile	Ser Leu Glu Arg 460 Ala Thr Glu His	Asp Phe Glu 445 Gly Cys Ala Asp Trp 525	Glu Val 430 Glu Pro Lys Glu Trp 510 Thr	Ser 415 Asn Ile Leu Ser Lys 495 Ile	400 Val Ser Gln Asn Leu 480 Asp Pro

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140 <211> LENGTH: 554 141 <212> TYPE: PRT 142 <213> ORGANISM: Rattus norvegicus 144 <220> FEATURE: 145 <223> OTHER INFORMATION: rat soluble epoxide hydrolase (sEH) 147 <400> SEQUENCE: 2 148 Met Ala Leu Arg Val Ala Ala Phe Asp Leu Asp Gly Val Leu Ala Leu 151 Pro Ser Ile Ala Gly Val Leu Arg His Thr Glu Glu Ala Leu Ala Leu 20 154 Pro Arg Asp Phe Leu Leu Gly Ala Phe Gln Met Lys Phe Pro Glu Gly 155 35 40 157 Pro Thr Glu Gln Leu Met Lys Gly Lys Ile Thr Phe Ser Gln Trp Val 158 50 55 160 Pro Leu Met Asp Glu Ser Cys Arg Lys Ser Ser Lys Ala Cys Gly Ala 70 163 Ser Leu Pro Glu Asn Phe Ser Ile Ser Glu Ile Phe Ser Gln Ala Met 90 166 Ala Ala Arg Ser Ile Asn Arg Pro Met Leu Gln Ala Ala Ala Leu 100 105 169 Lys Lys Gly Phe Thr Thr Cys Ile Val Thr Asn Asn Trp Leu Asp 115 120 125 172 Asp Ser Asp Lys Arg Asp Ile Leu Ala Gln Met Met Cys Glu Leu Ser 130 135 140 175 Gln His Phe Asp Phe Leu Ile Glu Ser Cys Gln Val Gly Met Ile Lys 150 155 178 Pro Glu Pro Gln Ile Tyr Lys Phe Val Leu Asp Thr Leu Lys Ala Lys 165 170 181 Pro Asn Glu Val Val Phe Leu Asp Asp Phe Gly Ser Asn Leu Lys Pro 180 185 184 Ala Arg Asp Met Gly Met Val Thr Ile Leu Val Arg Asp Thr Ala Ser 195 200 187 Ala Leu Arg Glu Leu Glu Lys Val Thr Gly Thr Gln Phe Pro Glu Ala 215 190 Pro Leu Pro Val Pro Cys Ser Pro Asn Asp Val Ser His Gly Tyr Val 230 235 193 Thr Val Lys Pro Gly Ile Arg Leu His Phe Val Glu Met Gly Ser Gly 245 250 196 Pro Ala Ile Cys Leu Cys His Gly Phe Pro Glu Ser Trp Phe Ser Trp 260 265 199 Arg Tyr Gln Ile Pro Ala Leu Ala Gln Ala Gly Phe Arg Val Leu Ala 275 280 202 Ile Asp Met Lys Gly Tyr Gly Asp Ser Ser Pro Pro Glu Ile Glu 295 300 205 Glu Tyr Ala Met Glu Leu Leu Cys Glu Glu Met Val Thr Phe Leu Asn 310 315 208 Lys Leu Gly Ile Pro Gln Ala Val Phe Ile Gly His Asp Trp Ala Gly 325 330 211 Val Leu Val Trp Asn Met Ala Leu Phe His Pro Glu Arg Val Arg Ala

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212
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 214 Val Ala Ser Leu Asn Thr Pro Leu Met Pro Pro Asn Pro Glu Val Ser
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 217 Pro Met Glu Val Ile Arg Ser Ile Pro Val Phe Asn Tyr Gln Leu Tyr
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 223 Arg Thr Phe Lys Ser Phe Phe Arg Thr Ser Asp Asp Met Gly Leu Leu
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 226 Thr Val Asn Lys Ala Thr Glu Met Gly Gly Ile Leu Val Gly Thr Pro
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 229 Glu Asp Pro Lys Val Ser Lys Ile Thr Thr Glu Glu Glu Ile Glu Tyr
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 232 Tyr Ile Gln Gln Phe Lys Lys Ser Gly Phe Arg Gly Pro Leu Asn Trp
 233 450
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238 Arg Lys Ile Leu Val Pro Ala Leu Met Val Thr Ala Glu Lys Asp Ile
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272 Pro Thr Glu Gln Leu Met Lys Gly Lys Ile Thr Phe Ser Gln Trp Val
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275 Pro Leu Met Asp Glu Ser Tyr Arg Lys Ser Ser Lys Ala Cys Gly Ala
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278 Asn Leu Pro Glu Asn Phe Ser Ile Ser Gln Ile Phe Ser Gln Ala Met
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281 Ala Ala Arg Ser Ile Asn Arg Pro Met Leu Gln Ala Ala Ile Ala Leu
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284 Lys Lys Gly Phe Thr Thr Cys Ile Val Thr Asn Asn Trp Leu Asp
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285			115					120					125			
		Glv			Ara	Asn	Ser			Gln	Mot	Mot			T ou	Ser
288		130		-7.5	3		135		. 1110	. OII	nec	140		Giu	. шеи	ser
290	Gln	His	Phe	Asn	Phe	Len			Ser	Cve	- Gln			Mot	Tla	T
291	145					150		Olu	. DCL	СуБ	155		GIŞ	Mec	ııe	_
		Glu	Pro	Gln	Tle			Dhe	T.011	Len			Ton	Tera	77-	160
294		o _z a		0111	165		Apii	FIIC	пец	170		1111	ьеи	г гув		гуѕ
		Asn	Glu	Val			T.011	Λαn	7 cn			Com	7	Т	175	ъ.,
297		11011	Olu	180		LIIC	пец	дад	185		Gry	ser	ASII		гÀг	Pro
		Arg	Agn			Mot	v-1	Thr			77-7	TT-1 ~	7	190	7 .7 -	a .
300		21129	195	MCC	Gry	Mec	vai	200	116	ьeu	val	HIS		Thr	Ala	ser
	Δla	Leu	-	Glu	T.011	Glu	T 770		The	~1	mbs	01 -	205	D	~ 1	
303	211.0	210	nr 9	Giu	пеα	Giu	215	val	1111	СТУ	1111		Pne	Pro	GIU	Ala
	Pro	Leu	Pro	17:⇒1	Dro	Cvc		Dro	7 02	7\ ~~	7707	220	TT-1	~ 1.		
	225		110	vai	FLO	230	ASII	PIO	ASII	Asp		ser	HIS	GLY	Tyr	
		Val	Lare	Dro	Clv		Λ×α	T 011	IIi a	Dha	235	α1	N/ - +	~ 1		240
309	1111	vai	цуз	FIO	245	116	Arg	ьец	птъ			GIU	мет	GIY		GIY
	Pro	Δ1=	T 11	Cvc		Crra	uic	C1.,	Dha	250 Dec		G		D1.	255	_
312	rio	Ala	шец	260	пеп	Cys	nis	GTA		Pro	GIU	ser	Trp		Ser	Trp
	Δra	ጥነታን	Gln		Dro	7/1 ~	T 011	77.	265	70 7 -	~1	D1	3	270	_	
315	Arg	Tyr	275	116	PIO	Ala	ьец		GIN	Ala	GIY	Pne		Val	Leu	Ala
	Tla	Λαn		Tara	C1	There	C7	280	0	Q	0	.	285	~3		
318	110	Asp 290	Mec	пуз	GIY	ıyı	295	Asp	ser	ser	ser		Pro	GIU	TTe	GIu
	Glu		715	Mot	C7.,,	T 011		C	T	a 1	1 4 - 1-	300	5 2.1	_,	_	_
	305	Tyr	AIA	Mec	GIU	310	ьец	Cys	ьуѕ	GIU		vaı	Thr	Phe	Leu	_
		T.e.11	Glv	בוד	Dro		ת ד ת	77-7	Dho	т1.	315	TT	7	m	2.7	320
324	пур	Leu	СТУ	116	325	GIII	Ala	Val	Pne		GIY	HIS	Asp	Trp		GLY
	Val	Met	Val	Trn		Mot	Nlα	Lou	Dho	330	Dro	a1	7	**- 7	335	77 -
327	Val	MCC	vai	340	ASII	Mec	нта	пец	345	TÀT	PIO	GIU	Arg		Arg	Ala
	Va1	Ala	Ser		Δen	Thr	Dro	Dho		Dro	Dwo	7	Dage	350	77 T	a
330			355	neu	11011	1111	110	360	Mec	PIO	PIO	Asp		Asp	vai	Ser
	Pro	Met		Val	Tle	Δra	Ser		Dro	Un l	Dho	7 00	365	~1 - -	T	Ш
333	~ = 0	370		vai	110	rrg	375	110	FIO	vai	FIIE	380	IÀT	GIII	ьеи	TYL
	Phe	Gln	Glu	Pro	Glv	Val	-	Glu.	Δla	Glu	T OU		T	7 000	Mot	C
336	385		024		OLY	390	nia	Olu	ALA	GIU	395	Gru	цуѕ	ASII	Met	
		Thr	Phe	Lvs	Ser		Dhe	Ara	בות	Sor		C1.,	Thr	~1··	Dha	400
339	9			<i></i> 1	405	1110	TIIC	лгg	Αια	410	Asp	GIU	TIII	GIY		шe
	Ala	Val	His	Lvs		Thr	Glu	Tle	Glv		Tla	Lou	\7.7.7	7 an	415	Dwo
342				420	1114		010	110	425	Gry	116	пец	val		1111	PIO
	Glu	Asp	Pro		T.eu	Ser	Lvc	Tla		Thr	Clu	C111	Clu	430	C1	Dha
345		L	435		204	DCI	275	440	1111	1111	GIU	GIU	445	116	GIU	Pile
	Tvr	Ile		Gln	Phe	Lvs	Lvs		Glv	Dhe	Λνα	G1 v		Ton	7 ~~	Пин
348	-1-	450	0111	0111	1110	1175	455	1111	Gry	FIIC	Arg	460	PLO	пец	ASII	тр
	Tvr	Arg	Δgn	Thr	Glu	Δrα		Trn	Larc	Trn	Car		T	~1	T	G1
351	465	5			Olu	470	Abii	TIP	шуз	rrb	475	Cys	nys	Gry	ьeu	
		Lys	Tle	Leu	Val		7\ T =	Lau	Mot	17-1		71 -	~1	T	7	480
354	5	-10		_ u	485		111 a	⊒Cu	171CL	490	TIII	нта	GIU	nys	_	тте
	Val	Leu	Ara	Pro		Met	Ser	Laze	λαν		C1	T 370	Т~~	т1 -	495	Dla -
357	, u.	⊥ cu	* 11 9	500	JIU	MEL	SET	пλр		Met	GIU	пув	ттр		rro	rne
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VERIFICATION SUMMARY

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